### WHAT IS CLAIMED IS:

### 1. A compound of Formula I

 $R^1$  N  $R^3$   $O-R^2$ 

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wherein

10 R<sup>1</sup> is selected from

- 1) hydrogen,
- 2) halogen,
- 3) substituted or unsubstituted C<sub>1</sub>-C<sub>10</sub> alkyl,
- 4) substituted or unsubstituted C2-C10 alkenyl,
- 5) substituted or unsubstituted C2-C10 alkynyl,
  - 6) substituted or unsubstituted aryl,
  - 7) substituted or unsubstituted C3-C10 cycloalkyl,
  - 8) substituted or unsubstituted heterocyclyl,
  - 9)  $-(CRa_2)_nOR^4$ , and

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10) –(CRa<sub>2</sub>)<sub>t</sub>C(O)OR<sup>4</sup>;

said alkyl, alkenyl, alkynyl, aryl, cycloalkyl, and heterocyclyl is optionally substituted with one or more of R<sup>7</sup>;

R<sup>2</sup> is selected from

- 25
- 1) hydrogen,
- 2) substituted or unsubstituted aralkyl,
- 3) substituted or unsubstituted C<sub>1</sub>-C<sub>10</sub> alkyl,
- 4) substituted or unsubstituted heterocyclyl,
- 5) substituted or unsubstituted aryl, and

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6) substituted or unsubstituted C3-C10 cycloalkyl;

# R<sup>3</sup> is selected from 1) hydrogen, 2) halogen, 3) $-C(O)R^4$ , 5 4) substituted or unsubstituted C1-C10 alkyl, substituted or unsubstituted aryl. 6) substituted or unsubstituted heterocyclyl, 7) substituted or unsubstituted C<sub>3</sub>-C<sub>10</sub> cycloalkyl, 8) substituted or unsubstituted C2-C10 alkenyl, and . 9) substituted or unsubstituted C2-C10 alkynyl; 10 R4 is independently selected from 1) hydrogen, substituted or unsubstituted C1-C10 alkyl, 15 3) substituted or unsubstituted aryl, 4) substituted or unsubstituted heterocyclyl, 5) substituted or unsubstituted C3-C10 cycloalkyl, 6) substituted or unsubstituted C2-C10 alkenyl, and 7) substituted or unsubstituted C2-C10 alkynyl; 20 R<sup>6</sup> is independently selected from 1) substituted or unsubstituted aryl, 2) substituted or unsubstituted heterocyclyl, 3) substituted or unsubstituted cycloalkyl, and 25 4) halogen; R<sup>7</sup> is independently selected from 1) hydrogen, 2) halogen, 30 3) substituted or unsubstituted C<sub>1</sub>-C<sub>10</sub> alkyl, 4) substituted or unsubstituted C2-C10 alkenyl,

5) substituted or unsubstituted C2-C10 alkynyl,
6) substituted or unsubstituted C3-C10 cycloalkyl,

7) substituted or unsubstituted aryl,

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8) substituted or unsubstituted heterocyclyl,
                       9) -NO<sub>2</sub>,
                       10) -NR_4(CRa_2)_nC(O)R^4,
                       11) -(CRa_2)<sub>n</sub>NR4_2,
                       12) -(CRa_2)_nNR^4(CRa_2)_nR^6,
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                       13) -CN,
                       14) -(CRa_2)_nC(O)R^4,
                       15) -(CRa_2)_nC(O)(CRa_2)_nOR^4,
                       16) -(CRa_2)_nOR^4,
                        17) –(CRa<sub>2</sub>)<sub>n</sub>R6,
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                        18) -(CRa_2)_nC(O)OR^4, and
                        19) -(CRa_2)_nSi(R^4)_3;
      Ra is independently selected from
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                        1) hydrogen,
                        2) substituted or unsubstituted C<sub>1</sub>-C<sub>10</sub> alkyl,
                        3) substituted or unsubstituted C2-C10 alkenyl,
                        4) substituted or unsubstitute C2-C10 alkynyl,
                        5) -OR4,
                        -C(O)OR^4,
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                        7) -NR^{4}_{2},
                        8) substituted or unsubstituted aryl,
                        9) substituted or unsubstituted heterocyclyl, and
                        10) substituted or unsubstituted C3-C10 cycloalkyl;
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       n is independently 0 to 6;
       t is 1 to 4;
       or a pharmaceutically acceptable salt or stereoisomer thereof.
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                        2.
                                 The compound according to Claim 1,
        wherein
        R1 is selected from
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                         1) hydrogen,
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- 2) halogen,
- 3) substituted or unsubstituted C<sub>1</sub>-C<sub>6</sub> alkyl,
- 4) substituted or unsubstituted C2-C10 alkynyl,
- 5) substituted or unsubstituted aryl,
- 6) substituted or unsubstituted C3-C10 cycloalkyl, and
- 7) substituted or unsubstituted heterocyclyl;

said alkyl, alkynyl, aryl, cycloalkyl, and heterocyclyl is optionally substituted with one or more of R<sup>7</sup>;

- 10 R<sup>2</sup> is selected from
  - 1) substituted or unsubstituted aralkyl,
  - 2) substituted or unsubstituted C1-C6 alkyl,
  - 3) substituted or unsubstituted aryl, and
  - 4) substituted or unsubstituted C3-C10 cycloalkyl;

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R<sup>3</sup> is selected from

- 1) halogen,
- 2)  $-C(O)R^4$ , and
- 3) substituted or unsubstituted C1-C6 alkyl;

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R<sup>4</sup> is independently selected from

- 1) hydrogen,
- substituted or unsubstituted C<sub>1</sub>-C<sub>6</sub> alkyl,
- 3) substituted or unsubstituted aryl,
- 4) substituted or unsubstituted heterocyclyl, and
- 5) substituted or unsubstituted C3-C10 cycloalkyl;

or a pharmaceutically acceptable salt or stereoisomer thereof.

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3. The compound according to Claim 2,

wherein

R<sup>1</sup> is selected from

- 1) substituted or unsubstituted C1-C6 alkyl,
- 35 2) substituted or unsubstituted C2-C10 alkynyl,

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- 3) substituted or unsubstituted heterocyclyl and
- 4) substituted or unsubstituted aryl;

said alkyl, alkynyl, heterocyclyl and aryl is optionally substituted with one or more of R7;

5 R<sup>2</sup> is selected from

- 1) substituted or unsubstituted aralkyl, and
- 2) substituted or unsubstituted C<sub>1</sub>-C<sub>6</sub> alkyl;

R<sup>3</sup> is selected from

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- 1) halogen, and
- 2)  $-C(O)R^4$ ;

or a pharmaceutically acceptable salt or stereoisomer thereof.

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4. A compound of Formula II

$$R^1$$
 $N$ 
 $H$ 
 $O$ 
 $O-R^2$ 
 $II$ 

wherein

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R1 is selected from

- 1) hydrogen,
- 2) halogen,
- 3) substituted or unsubstituted C<sub>1</sub>-C<sub>6</sub> alkyl,

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- 4) substituted or unsubstituted C2-C10 alkynyl,
- 5) substituted or unsubstituted aryl,
- 6) substituted or unsubstituted C3-C10 cycloalkyl, and
- 7) substituted or unsubstituted heterocyclyl;

said alkyl, alkynyl, aryl, cycloalkyl and heterocyclyl is optionally substituted with one or more of R<sup>7</sup>;

#### R<sup>2</sup> is selected from

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- 1) substituted or unsubstituted aralkyl, and
- 2) substituted or unsubstituted C<sub>1</sub>-C<sub>6</sub> alkyl;
- 5 R4 is independently selected from
  - 1) hydrogen,
  - 2) substituted or unsubstituted C1-C10 alkyl,
  - 3) substituted or unsubstituted aryl,
  - 4) substituted or unsubstituted heterocyclyl,
  - 5) substituted or unsubstituted C3-C10 cycloalkyl,
  - 6) substituted or unsubstituted C2-C10 alkenyl, and
  - 7) substituted or unsubstituted C2-C10 alkynyl;

## R<sup>6</sup> is independently selected from

- 1) substituted or unsubstituted aryl,
  - 2) substituted or unsubstituted heterocyclyl,
  - 3) substituted or unsubstituted C3-C10 cycloalkyl, and
  - 4) halogen;
- 20 R7 is independently selected from
  - 1) hydrogen,
  - 2) halogen,
  - 3) substituted or unsubstituted C<sub>1</sub>-C<sub>10</sub> alkyl,
  - 4) substituted or unsubstituted C2-C10 alkenyl,
  - 5) substituted or unsubstituted C2-C10 alkynyl,
  - 6) substituted or unsubstituted C3-C10 cycloalkyl,
  - 7) substituted or unsubstituted aryl,
  - 8) substituted or unsubstituted heterocyclyl,
  - 9) -NO<sub>2</sub>,
- 30 10)  $-NR^4(CRa_2)_nC(O)R^4$ ,
  - 11) -( $CRa_2$ )<sub>n</sub> $NR4_2$ ,
  - 12)  $-(CRa_2)_nNR^4(CRa_2)_nR^6$ ,
  - 13) -CN,
  - 14) - $(CR^{a}_{2})_{n}C(O)R^{4}$ ,

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15) -(CRa_2)_nC(O)(CRa_2)_nOR4,
                        16) -(CRa_2)_nOR4,
                        17) -(CRa_2)_nR6,
                        18) -(CRa_2)_nC(O)OR4, and
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                        19) -(CRa_2)_nSi(R^4)_3;
      Ra is independently selected from
                       1) hydrogen,
                       2) substituted or unsubstituted C_1-C_{10} alkyl,
                       3) substituted or unsubstituted C<sub>1</sub>-C<sub>10</sub> alkenyl,
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                       4) substituted or unsubstitute C<sub>1</sub>-C<sub>10</sub> alkynyl,
                       5) -OR4,
                       -C(O)OR4
                       7) -NR^{4}_{2},
15
                       8) substituted or unsubstituted aryl,
                       9) substituted or unsubstituted heterocyclyl, and
                       10) substituted or unsubstituted C3-C10 cycloalkyl;
      n is independently 0 to 6;
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      t is 1 to 4;
      or a pharmaceutically acceptable salt or stereoisomer thereof.
                       5.
                               A compound selected from:
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      benzyl 4-ethyl-2-formyl-1H-pyrrole-3-carboxylate;
      benzyl 4-ethyl-2-formyl-5-iodo-1H-pyrrole-3-carboxylate;
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      methyl 4-ethyl-2-formyl-5-iodo-1H-pyrrole-3-carboxylate;
      methyl 4-ethyl-2,5-diiodo-1H-pyrrole-3-carboxylate;
      methyl 5-(4-fluorophenyl)-4-ethyl-2-formyl-1H-pyrrole-3-carboxylate;
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methyl 4-ethyl-2-formyl-5-thien-2-yl-1H-pyrrole-3-carboxylate;
      methyl 4-ethyl-2-formyl-5-[3-(trimethylsilyl)prop-1-ynyl]-1H-pyrrole-3-carboxylate;
 5
      4'-benzyl 1-tert-butyl 3'-ethyl-5'-formyl-1H,1'H-2,2'-bipyrrole-1,4'-dicarboxylate;
      benzyl 5-(3,5-dimethylisoxazol-4-yl)-4-ethyl-2-formyl-1H-pyrrole-3-carboxylate;
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      benzyl 5-(1-benzofuran-2-yl)-4-ethyl-2-formyl-1H-pyrrole-3-carboxylate;
      benzyl 4-ethyl-2-formyl-5-(3-nitrophenyl)-1H-pyrrole-3-carboxylate;
      benzyl 4-ethyl-2-formyl-5-(5-methyl-2-furyl)-1H-pyrrole-3-carboxylate;
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      benzyl 5-[3-(acetylamino)phenyl]-4-ethyl-2-formyl-1H-pyrrole-3-carboxylate;
      benzyl 4-ethyl-2-formyl-5-pyridin-4-yl-1H-pyrrole-3-carboxylate;
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      benzyl 4-ethyl-2-formyl-5-phenyl-1H-pyrrole-3-carboxylate;
      benzyl 5-(3-cyanophenyl)-4-ethyl-2-formyl-1H-pyrrole-3-carboxylate;
      benzyl 4-ethyl-2-formyl-5-(3-methoxyphenyl)-1H-pyrrole-3-carboxylate;
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      benzyl 4-ethyl-2-formyl-5-(5-formyl-2-furyl)-1H-pyrrole-3-carboxylate:
      methyl 4-ethyl-2-formyl-5-(phenylethynyl)-1H-pyrrole-3-carboxylate;
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      methyl 5-{3-[benzyl(methyl)amino]prop-1-ynyl}-4-ethyl-2-formyl-1H-pyrrole-3-carboxylate;
      benzyl 5-(2-cyanophenyl)-4-ethyl-2-formyl-1H-pyrrole-3-carboxylate;
      benzyl 5-(4-cyanophenyl)-4-ethyl-2-formyl-1H-pyrrole-3-carboxylate;
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benzyl 4-ethyl-2-formyl-5-(4-nitrophenyl)-1H-pyrrole-3-carboxylate;
       benzyl 4-ethyl-2-formyl-5-(2-methoxyphenyl)-1H-pyrrole-3-carboxylate;
       benzyl 4-ethyl-2-formyl-5-(4-methoxyphenyl)-1H-pyrrole-3-carboxylate;
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       benzyl 4-ethyl-2-formyl-5-(2-methylphenyl)-1H-pyrrole-3-carboxylate;
       benzyl 4-ethyl-2-formyl-5-(3-methylphenyl)-1H-pyrrole-3-carboxylate;
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       benzyl 5-(2-chlorophenyl)-4-ethyl-2-formyl-1H-pyrrole-3-carboxylate;
       benzyl 5-(3-chlorophenyl)-4-ethyl-2-formyl-1H-pyrrole-3-carboxylate;
       methyl 4-ethyl-2-formyl-5-[1-(3-hydroxypropyl)vinyl]-1H-pyrrole-3-carboxylate;
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       methyl 4-ethyl-2-formyl-5-(5-hydroxypent-1-ynyl)-1H-pyrrole-3-carboxylate;
       methyl 4-ethyl-2-formyl-5-[(1-hydroxycyclohexyl)ethynyl]-1H-pyrrole-3-carboxylate;
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       methyl 5-[3-(dimethylamino)prop-1-ynyl]-4-ethyl-2-formyl-1H-pyrrole-3-carboxylate;
       methyl 5-(3,3-dimethylbut-1-ynyl)-4-ethyl-2-formyl-1H-pyrrole-3-carboxylate;
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       methyl 4-ethyl-2-formyl-5-(pyridin-2-ylethynyl)-1H-pyrrole-3-carboxylate;
       methyl 4-ethyl-2-formyl-5-(6-methoxypyridin-2-yl)-1H-pyrrole-3-carboxylate;
      methyl 4-ethyl-2-formyl-5-(3-methoxyprop-1-ynyl)-1H-pyrrole-3-carboxylate;
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      methyl\ 5\hbox{--}[(2\hbox{-bromophenyl})\hbox{-ethyl-}2\hbox{-formyl-}1\hbox{H-pyrrole-}3\hbox{-carboxylate};
      methyl\ 5\hbox{-}[3\hbox{-}(1H\hbox{-}1,2,3\hbox{-}benzotriazol\hbox{-}1\hbox{-}yl)prop-1\hbox{-}ynyl]\hbox{-}4\hbox{-}ethyl\hbox{-}2\hbox{-}formyl\hbox{-}1H\hbox{-}pyrrole\hbox{-}3\hbox{-}pyrolayl)prop-1\hbox{-}ynyl]}
       carboxylate;
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methyl 4-ethyl-2-formyl-5-(4-methylpyridin-2-yl)-1H-pyrrole-3-carboxylate;
      methyl 4-ethyl-2-formyl-5-(6-methylpyridin-2-yl)-1H-pyrrole-3-carboxylate;
 5
      methyl 5-(4-tert-butylphenyl)-4-ethyl-2-formyl-1H-pyrrole-3-carboxylate;
      methyl 5-(2,4-difluorophenyl)-4-ethyl-2-formyl-1H-pyrrole-3-carboxylate;
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      methyl 4-ethyl-2-formyl-5-[3-(methoxycarbonyl)phenyl]-1H-pyrrole-3-carboxylate;
      methyl 4-ethyl-2-formyl-5-[4-(methoxycarbonyl)phenyl]-1H-pyrrole-3-carboxylate;
      methyl 4-ethyl-2-formyl-5-[(1-hydroxycyclopentyl)ethynyl]-1H-pyrrole-3-carboxylate;
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      methyl 4-ethyl-2-formyl-5-(3-hydroxy-3-methylbut-1-ynyl)-1H-pyrrole-3-carboxylate
      methyl 4-ethyl-2-formyl-5-(1-hexylvinyl)-1H-pyrrole-3-carboxylate;
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      methyl 4-ethyl-2-formyl-5-(1,3-thiazol-2-yl)-1H-pyrrole-3-carboxylate;
      methyl 5-[1-(3-chloropropyl)vinyl]-4-ethyl-2-formyl-1H-pyrrole-3-carboxylate;
      methyl 5-(5-chloropent-1-ynyl)-4-ethyl-2-formyl-1H-pyrrole-3-carboxylate;
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      methyl 4-ethyl-2-formyl-5-(3-hydroxy-3-phenylbut-1-ynyl)-1H-pyrrole-3-carboxylate;
      methyl 4-ethyl-2-formyl-5-(3-methylpyridin-2-yl)-1H-pyrrole-3-carboxylate;
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      methyl 4-ethyl-2-formyl-5-isopentyl-1H-pyrrole-3-carboxylate;
      methyl 4-ethyl-2-formyl-5-(3-methylthien-2-yl)-1H-pyrrole-3-carboxylate:
      methyl 4-ethyl-2-formyl-5-isobutyl-1H-pyrrole-3-carboxylate;
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methyl 5-cyclohexyl-4-ethyl-2-formyl-1H-pyrrole-3-carboxylate;

methyl 5-butyl-4-ethyl-2-formyl-1H-pyrrole-3-carboxylate;

5 methyl 5-cyclopentyl-4-ethyl-2-formyl-1H-pyrrole-3-carboxylate;

methyl 5-(cyclohexylmethyl)-4-ethyl-2-formyl-1H-pyrrole-3-carboxylate;

methyl 5-sec-butyl-4-ethyl-2-formyl-1H-pyrrole-3-carboxylate;

methyl 4-ethyl-2-formyl-5-(3-methoxy-2-methyl-3-oxopropyl)-1H-pyrrole-3-carboxylate;

methyl 4-ethyl-2-formyl-5-phenyl-1H-pyrrole-3-carboxylate;

methyl 4-ethyl-2-formyl-5-pyridin-4-yl-1H-pyrrole-3-carboxylate;

methyl 4-ethyl-2-formyl-5-(4-nitrophenyl)-1H-pyrrole-3-carboxylate;

methyl 4-ethyl-2-formyl-5-(2-methoxyphenyl)-1H-pyrrole-3-carboxylate;

or a pharmaceutically acceptable salt or stereoisomer thereof.

- 6. The compound according Claim 5 that is selected from
- 25 methyl 4-ethyl-2-formyl-5-iodo-1H-pyrrole-3-carboxylate

benzyl 4-ethyl-2-formyl-5-phenyl-1H-pyrrole-3-carboxylate

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methyl 4-ethyl-2-formyl-5-[(1-hydroxycyclohexyl)ethynyl]-1H-pyrrole-3-carboxylate

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 $methyl\ 4-ethyl-2-formyl-5-(6-methoxypyridin-2-yl)-1 H-pyrrole-3-carboxylate$ 

methyl 5-[1-(3-chloropropyl)vinyl]-4-ethyl-2-formyl-1H-pyrrole-3-carboxylate

or a pharmaceutically acceptable salt or stereoisomer thereof.

7. A trifluoroacetic acid salt of a compound of Claim 5 which is selected from methyl 4-ethyl-2-formyl-5-(6-methoxypyridin-2-yl)-1H-pyrrole-3-carboxylate; 5 methyl 4-ethyl-2-formyl-5-(4-methylpyridin-2-yl)-1H-pyrrole-3-carboxylate; methyl 4-ethyl-2-formyl-5-(6-methylpyridin-2-yl)-1H-pyrrole-3-carboxylate; 10 benzyl 4-ethyl-2-formyl-5-pyridin-4-yl-1H-pyrrole-3-carboxylate. 8. A pharmaceutical composition which is comprised of a compound in accordance with Claim 1 and a pharmaceutically acceptable carrier. 15 9. A method of modulating the catalytic activity of protein kinases in a mammal in need thereof comprising contacting the protein kinase with a compound of Claim 1. 10. The method of Claim 9 wherein the protein kinase is an RTK. 20 11. The method of Claim 10, wherein the RTK is selected from IR, IGF-1R and IRR. 12. A method of treating a PK-related disorder in a mammal in need 25 thereof comprising administering to said mammal a therapeutically effective amount of a compound of Claim 1. 13. A method of Claim 12, wherein the PK-related disorder is an IGF-1R-related disorder selected from: 30 1) cancer, 2) diabetes. 3) an autoimmune disorder,

a hyperproliferation disorder,

4)

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6)

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aging,

acromegaly, and

- 7) Crohn's disease.
- 14. A method of preventing a PK-related disorder in a mammal in need thereof comprising administering to said mammal a therapeutically effective amount of a compound of Claim 1.
  - 15. A method of Claim 14, wherein the PK-related disorder is an IGF-1R-related disorder selected from:
    - 1) cancer,
- 10 2) diabetes,
  - 3) an autoimmune disorder,
  - 4) a hyperproliferation disorder,
  - 5) aging,
  - 6) acromegaly, and
- 15 7) Crohn's disease.
  - 16. A method of treating cancer in a mammal in need of such treatment comprising administering to said mammal a therapeutically effective amount of a compound of Claim 1.

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- 17. A method of treating retinal vascularization comprising administering to a mammal in need of such treatment a therapeutically effective amount of a compound of Claim 1.
- 25 18. A method of treating cancer which comprises administering a therapeutically effective amount of a compound of Claim 1 in combination with a second compound selected from:
  - 1) an estrogen receptor modulator,
  - 2) an androgen receptor modulator,
  - 3) a retinoid receptor modulator,
  - 4) a cytotoxic/cytostatic agent,
  - 5) an antiproliferative agent,
  - 6) a prenyl-protein transferase inhibitor,
  - 7) an HMG-CoA reductase inhibitor,
- 35 an HIV protease inhibitor,

- 9) a reverse transcriptase inhibitor,
- 10) an angiogenesis inhibitor,
- 11) a PPAR-y agonist,
- 12) a PPAR-δ agonists,
- 13) an inhibitor of cell proliferation and survival signaling, and
- 14) an agent that interferes with a cell cycle checkpoint.
- 19. The method of Claim 18, wherein the second compound is an estrogen receptor modulator selected from tamoxifen and raloxifene.

20. A method of preventing cancer which comprises administering a therapeutically effective amount of a compound of Claim 1 in combination with a second compound selected from:

- 1) an estrogen receptor modulator,
- 2) an androgen receptor modulator,
- 3) a retinoid receptor modulator,
- 4) a cytotoxic/cytostatic agent,
- 5) an antiproliferative agent,
- 6) a prenyl-protein transferase inhibitor,
- 7) an HMG-CoA reductase inhibitor,
- 8) an HIV protease inhibitor,
- 9) a reverse transcriptase inhibitor,
- 10) an angiogenesis inhibitor,
- 11) a PPAR-y agonist,
- 12) a PPAR-δ agonists,
  - 13) an inhibitor of cell proliferation and survival signaling, and
  - 14) an agent that interferes with a cell cycle checkpoint.
- 21. The method of Claim 20, wherein the second compound is an estrogen receptor modulator selected from tamoxifen and raloxifene.
  - 22. A method of treating cancer which comprises administering a therapeutically effective amount of a compound of Claim 1 in combination with radiation therapy.

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23. The method of Claim 22 wherein radiation therapy is also administered.

- A method of treating cancer which comprises administering a
   therapeutically effective amount of a compound of Claim 1 and paclitaxel or trastuzumab.
  - 25. A method of treating cancer which comprises administering a therapeutically effective amount of a compound of Claim 1 and a GPIIb/IIIa antagonist.
- 10 26. The method of Claim 26 wherein the GPIIb/IIIa antagonist is tirofiban.
  - 27. A method of preventing cancer which comprises administering a therapeutically effective amount of a compound of Claim 1 and a GPIIb/IIIa antagonist.
  - 28. The method of Claim 27 wherein the GPIIb/IIIa antagonist is tirofiban.
- 29. A method of treating cancer which comprises administering a therapeutically effective amount of a compound of Claim 1 in combination with a COX-2 inhibitor.
- 30. A method of preventing cancer which comprises administering a therapeutically effective amount of a compound of Claim 1 in combination with a COX-2 inhibitor.

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